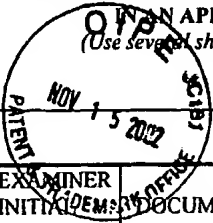


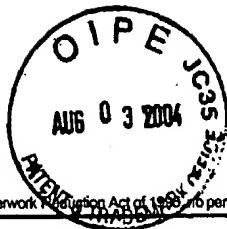
Form PTO/SB/08 EXAMINATION DISCLOSURE CITATION IN AN APPLICATION (Use several sheets if necessary)		Docket Number (Optional) CIBT-P10-203		Application Number 09/754,032		
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		Filing Date January 3, 2001				Group Art Unit 1646
U.S. PATENT DOCUMENTS						
EXAMINER INITIAL	DOCUMENT NUMBER	DATE	NAME	CLASS	SUBCLASS	FILING DATE IF APPROPRIATE
FOREIGN PATENT DOCUMENTS						TECH CENTER 1600/2900
	DOCUMENT NUMBER	DATE	COUNTRY	CLASS	SUBCLASS	Translation YES NO
OTHER DOCUMENTS (Including Author, Title, Date, Pertinent Pages Etc.)						
<div style="font-size: 2em;"> DT ↓ </div>	DT	Hammerschmidt, M. et al. Protein kinase A is a common negative regulator of Hedgehog signaling in the vertebrate embryo. <i>Genes & Development</i> 10, 647-658 (1996).				
<div style="font-size: 2em;"> DU ↓ </div>	DU	Noveen, A. et al. cAMP, an Activator of Protein Kinase A, Suppresses the Expression of Sonic Hedgehog. <i>Biochem. Biophys. Res. Comm.</i> 219, 180-185 (1996).				
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		Applicant Scott et al.		Group Art Unit 1646			
		Filing Date January 3, 2001					
U.S. PATENT DOCUMENTS							
EXAMINER INITIALS	DOCUMENT NUMBER	DATE	NAME	CLASS	FILING DATE IF APPROPRIATE		
<i>W</i>	DV 6,027,882	2/22/00	Scott et al.				
FOREIGN PATENT DOCUMENTS							
	DOCUMENT NUMBER	DATE	COUNTRY	CLASS	SUBCLASS	Translation YES NO	
OTHER DOCUMENTS						(Including Author, Title, Date, Pertinent Pages Etc.)	
<i>W</i>	DW	Mullins, L.J. & Mullins, J.J. Transgenesis in the Rat and Larger Mammals. <i>J. Clin. Invest.</i> 98, S37-S40 (1996).					
	DX	Ngo, J.T. et al. Computational Complexity, Protein Structure Prediction, and the Levinthal Paradox. <i>Peptide Hormones</i> 491-494 (1976).					
	DY	Rudinger, J. Characteristics of the amino acids as components of a peptide hormone sequence. <i>Endocrinology</i> 5-7 (1972).					
	DZ	Seamark, R.F. Progress and Emerging Problems in Livestock Transgenesis: a Summary Perspective. <i>Reprod. Fert. Dev.</i> 6, 653-657 (1994).					
<i>W</i>	EA	Wall, R.J. Transgenic Livestock: Progress and Prospects for the Future. <i>Theriogenology</i> 45, 57-68 (1996).					
EXAMINER			DATE CONSIDERED				
<i>[Signature]</i>			9/3/04				
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Substitute for form 1449A/B/PTO INFORMATION DISCLOSURE STATEMENT BY APPLICANT (Use as many sheets as necessary)				Complete if Known	
				Application Number	09/754032
				Filing Date	January 3, 2001
				First Named Inventor	Matthew P. Scott
				Art Unit	1646
				Examiner Name	M. T. Brannock
Sheet	1	of	1	Attorney Docket Number	CIBT-P10-203

U.S. PATENT DOCUMENTS					
Examiner Initials*	Cite No. ¹	Document Number	Publication Date MM-DD-YYYY	Name of Patentee or Applicant of Cited Document	Pages, Columns, Lines, Where Relevant Passages or Relevant Figures Appear
		Number-Kind Code ² (if known)			

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Examiner Initials*	Cite No. ¹	Foreign Patent Document	Publication Date MM-DD-YYYY	Name of Patentee or Applicant of Cited Document	Pages, Columns, Lines, Where Relevant Passages or Relevant Figures Appear	T ⁶
		Country Code ³ -Number ⁴ -Kind Code ⁵ (if known)				

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Examiner Initials*	Cite No. ¹	Include name of the author (in CAPITAL LETTERS), title of the article (when appropriate), title of the item (book, magazine, journal, serial, symposium, catalog, etc.), date, page(s), volume-issue number(s), publisher, city and/or country where published.				T ²
	CA	ALTABA et al., Cell 90:193-196, 1997				
	CB	FAN et al., Cell 81(3): 457-465, 5/5/1995				
	CC	KRISHNAN et al., Science 278: 1947-1950, 12/12/97				
	CD	MULLER et al., Development 127: 2999-3007, 2000				

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¹Applicant's unique citation designation number (optional). ²Applicant is to place a check mark here if English language Translation is attached.

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Docket Number (Optional)
CIBT-P10-203

Application Number
09/754,032

Applicant
Scott et al.

Filing Date
03-Jan-2001

Group Art. No.
1646

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W/PATENT EA	5798209	25-Aug-1998	Chan	435	6	26-May-1995
AB	5837538	17-Nov-1998	Scott	435	325	06-Oct-1995
AC	5935810	10-Aug-1999	Friedman et al.	435	69.1	30-Nov-1994

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						YES	NO
W	AD	WO9611260	18-Apr-1996	PCT	C12N	5/00	

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W	AE	Akimaru, H. et al., "Drosophila CBP is a co-activator of cubitus interruptus in hedgehog signaling", Nature 386 (6626): 735-738 (1997).
	AF	Akiyama, H. et al., "Cloning of a mouse smoothened cDNA and expression patterns of hedgehog signaling molecules during chondrogenesis and cartilage differentiation in conal mouse EC cells, ATDC5", Biophys Res. Comm., 235(1): 142-147 (1997).
	AG	Alberts, eds., Molecular Biology of the Cell, G-10 (1994).
	AH	Alcedo, J. et al., "The Drosophila smoothened gene encodes a seven-pass membrane protein, a putative receptor for the hedgehog signal", Cell, 86 (2): 221-232 (1996).
	AI	Alcedo, J. and Noll, M., "Hedgehog and its patched-smoothened receptor complex: a novel signaling mechanism at the cell surface", Biol. Chem., 378 (7): 583-590 (1997).
	AJ	Alexandre, C. et al., "Transcriptional activation of hedgehog target genes in Drosophila is mediated directly by the cutitus interruptus protein, a member of the GLI family of zinc finger DNA-binding proteins", Genes Dev., 19 (16): 2003-2013 (1996).
	AK	Bale, A., "Variable expressivity of patched mutations in flies and humans", Am. J. Human Genet., 60 (1): 10-12 (1997).
	AL	Bellusci, S. et al., "Involvement of Sonic hedgehog (Shh) in mouse embryonic lung growth and morphogenesis", Development, 124 (1): 53-63 (1997).
	AM	Bhat, K. and Schedl, P., "Requirement for engrailed and invected genes reveals novel regulatory interactions between engrailed/invected, patched, gooseberry and wingless during Drosophila neurogenesis", Development, 124 (9): 1675-1688 (1997).
	AN	Bitgood, M. et al., "Sertoli cell signaling by Desert hedgehog regulates the male germline", Curr. Biol., 6 (3): 298-304 (1996).
	AO	Bokor, P. et al., "The roles of hedgehog, wingless and lines in patterning the dorsal epidermis in Drosophila", Development, 122 (4): 1083-1092 (1996).
	AP	Bowie et al., "Deciphering the Message in Protein Sequences: Tolerance to Amino Acid Substitutions", Science 247: 1306-1310 (1990).

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INFORMATION DISCLOSURE CITATION IN AN APPLICATION (attach several sheets if necessary)		Applicant Scott et al.	
		Filing Date 03-Jan-2001	Group Art Unit 1646
O I P E AUG 31 2001 PATENT & TRADEMARK JC138	AO	Cadigan, K. et al., "Localized expression of sloppy paired protein maintains the polarity of Drosophila parasegments", Genes Dev., 8 (8): 899-913 (1994).	
	AR	Chanut, F. and Heberlein, U., "Role of the morphogenetic furrow in establishing polarity in the Drosophila eye", Development, 121 (12): 4085-1094 (1995).	
	AS	Chavrier et al., "The complexity of the Rab and Rho GTP-binding protein subfamilies revealed by a PCR cloning approach", Gene 112: 261-264 (1992).	
	AT	Chen, E. et al., "Compartmental organization of the Drosophila genital imaginal disks", Development, 124 (1): 205-218 (1997).	
	AU	Chen, Y. et al., "Dual roles for patched in sequestering and transducing Hedgehog", Cell, 87(3): 553-563 (1996).	
	AV	Concordet, J. et al., "Spatial regulation of a zebrafish patched homologue reflects the roles of sonic hedgehog and protein kinase A in neural tube and somite patterning", Development, 122 (9): 2835-2846 (1996).	
	AW	Dhawan et al., "Systematic Delivery of Human Growth Hormone by Injection of Genetically Engineered Myoblasts", Science 254: 1509-1512 (1991).	
	AX	Dominguez, M. et al., "Sending and receiving the hedgehog signal: control by the Drosophila Gli protein cubitus interruptus", Science, 272 (5268): 1621-1625 (1996).	
	AY	Echelard, Y. et al., "Sonic hedgehog, a member of a family of putative signaling molecules, is implicated in the regulation of CNS polarity", Cell, 75: 1417-1430 (1993).	
	AY	Epps, J. et al., "Orshigane, a new segment polarity gene of Drosophila melanogaster, functions in hedgehog signal transduction", Genetics 145 (4): 1041-1052 (1997).	
	BA	Epstein, D. et al., "Antagonizing cAMP-dependent protein kinase A in the dorsal CNS activates a conserved Sonic hedgehog signaling pathway", Development, 122 (9): 2884-2894 (1996).	
	BB	Forbes, A. et al., "The role of segment polarity genes during early oogenesis in Drosophila", Development, 122 (10): 33283-3294 (1996).	
	BC	Gailani et al., "Developmental Genes and Cancer: Role of Patched in Basal Cell Carcinoma of the Skin", J. Nat. Canc. Inst. 89 (15): 1103-1109 (1997).	
	BD	Gailani, M. et al., "The role of the human homologue of Drosophila patched in spradic basal cell carcinomas", Nat. Genet., 14 (1): 78-81 (1996).	
	BE	Gomez-Skarmeta, J.L. et al., "Araucan and caupolican provide a link between compartment subdivisions and patterning of sensory organs and veins in the Drosophila wing", Genes Dev., 10 (22): 2935-1945 (1996).	
	BF	Goodrich, L. et al., "Altered neural cell fates and medulloblastoma in mouse patched mutants", Science, 277 (5329): 1109-1113 (1997).	
BG	Goodrich, L. et al., "Conservation of the hedgehog/patched signaling pathway from flies to mice: induction of a mouse patched gene by Hedgehog", Genes Dev., 10 (3): 301-312 (1996).		

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		Gorlin, R., "Nevoid basal-cell carcinoma syndrome", Medicine, 66: 98-113 (1987).	
		Grindley, J. et al., "Evidence for the involvement of the Gli gene family in embryonic mouse lung development", Dev. Biol., 188 (2): 337-348 (1997).	
	BH	Habuchi, et al., "Detailed deletion mapping of chromosome 9q bladder cancer: evidence of two tumour suppressor loci", Oncogene, 11:1671-1674 (1995).	
	BJ	Hahn, H. et al., "A mammalian patched homolog is expressed in target tissues of sonic hedgehog and maps to a region associated with development abnormalities", J. Biol. Chem., 271 (21): 12125-12128 (1996).	
	BK	Heemskerk, J. et al., "Drosophila hedgehog acts as a morphogen in cellular patterning", Cell 76: 449-460 (1994).	
	BL	Hepker, J. et al., "Drosophila cubitus interruptus forms a negative feedback loop with patched and regulates expression of Hedgehog target genes", Development, 124 (2): 549-558 (1997).	
	BM	Hidalgo, A. and Ingham, P., "Cell patterning in the Drosophila segment: spatial regulation of the segment polarity gene patched", Development, 110: 291-301 (1990).	
	BN	Hooper et al., "The Drosophila patched gene encodes a putative membrane protein required for segmental patterning", Cell 59: 751-765 (1989).	
	BO	Hynes, M., et al., "Control of cell pattern in the neural tube by zinc finger transcription factor and oncogene _____ Neuron 19(1): 1997).	
	BP	Ingham, "Hedgehog points the way", Curr. Biol. 4: 347-350 (1994).	
	BQ	Ingham, P. et al., "Role of the Drosophila patched gene in positional signalling", Nature, 353: 184-187 (1991).	
	BR	Ingham, P. et al., "Quantitative effects of hedgehog and decapentaplegic activity on the patterning of the Drosophila wing", Curr. Biol., 5 (4): 432-440 (1995).	
	BS	Jensen, A. et al., "Expression of Sonic hedgehog and its putative role as a precursor cell mitogen in the developing mouse retina", Development, 124 (2): 363-371 (1997).	
	BT	Jiang, J. et al., "Protein kinase A and hedgehog signaling in Drosophila limb development", Cell, 80 (4): 563-572 (1995).	
	BU	Johnson, R. et al., "Patched overexpression alters wing disc size and pattern: transcriptional and post-transcriptional effects on hedgehog targets", Development, 121 (12): 4161-4170 (1995).	
	BV	Johnson, R. et al., "Human homolog of patched, a candidate gene for the basal cell nevus syndrome", Science, 272 (5268): 1668-1671 (1996).	
	BW	Kalderon, D., "Morphogenetic signalling. Responses to hedgehog", Curr. Biol., 5 (6): 2279-2289 (1995).	
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		Filing Date 03-Jan-2001	Group Art Unit 1646
Ito, Nippon Yakurigaku Zasshi, Folia Pharmacologica Japonica 102 (3): AN 94010590.			
	BY	Kojima, T. et al., "Induction of a mirror-image duplication of anterior wing structures by localized hedgehog expression in the anterior compartment of <i>Drosophila melanogaster</i> wing imaginal discs", <i>Gene</i> , 148 (2): 211-7 (1994).	
	CA	Krauss, S. et al., "A functionally conserved homolog of the <i>Drosophila</i> segment polarity gene <i>hh</i> is expressed in tissues with polarizing activity in zebrafish embryos", <i>Cell</i> , 75: 1431-1444 (1993).	
	CB	Lepage et al., "Signal transduction by cAMP-dependent protein kinase A in <i>Drosophila</i> limb patterning", <i>Nature</i> , 373 (6516): 711-715 (1995).	
	CC	Li, W., et al., "Function of protein kinase A in hedgehog signal transduction and <i>Drosophila</i> imaginal disc development", <i>Cell</i> , 80 (4): 553-562 (1995).	
	CD	Loftus, S., et al., "Murine model of Niemann-Pick C disease: mutation in a cholesterol homeostatis gene", <i>Science</i> , 277 (5323): 232-235 (1997).	
	CE	Ma, C. et al., "The segment polarity gene hedgehog is required for progression of the morphogenic furrow in the developing <i>Drosophila</i> eye", <i>Cell</i> , 75 (5): 927-938 (1993).	
	CF	Ma, C. et al., "Wingless and patched are negative regulators of the morphogenetic furrow and can effect tissue polarity in the developing <i>Drosophila</i> compound eye", <i>Development</i> , 121 (8): 2279-2289 (1995).	
	CG	Ma et al., "Molecular Cloning and Characterization of rKik10, a cDNA encoding T-Kininogenase from Rat Submandibular Gland and Kidney", <i>Biochemistry</i> 31: 10922-10928.	
	CH	Marigo, V. et al., "Biochemical evidence that patched is the Hedgehog receptor", <i>Nature</i> , 384 (6605): 176-179 (1996).	
	CI	Marigo, V. et al., "Conservation in hedgehog signaling: induction of a chicken patched homolog by Sonic hedgehog in the developing limb", <i>Development</i> , 122 (4): 1225-1233 (1996).	
	CJ	Marigo, V. et al., "Sonic hedgehog differentially regulates expression of GLI and GLI3 during limb development", <i>Dev. Biol.</i> , 180 (1): 273-283 (1996).	
	CK	Marigo, V. et al., "Regulation of patched by sonic hedgehog in the developing neural tube", <i>Proc. Natl. Acad. Sci. USA</i> , 93(18): 9346-9351 (1996).	
	CL	Nakamura, T. et al., "Induction of osteogenic differentiation by hedgehog proteins", <i>Biochem. Biophys. Res. Comm.</i> , 237 (2): 465-469 (1997).	
	CM	Nakano, Y. et al., "A protein with several possible membrane-spanning domains encoded by the <i>Drosophila</i> segment polarity gene patched", <i>Nature</i> , 341: 508-513 (1989).	
	CN	Nusse, R. "Patching up Hedgehog", <i>Nature</i> , 384 (6605): 119-120 (1996).	
	CO	Oro, A. et al., "Basal cell carcinomas in mice overexpressing sonic hedgehog", <i>Science</i> , 276(5313): 817-821 (1997).	

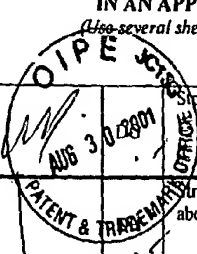
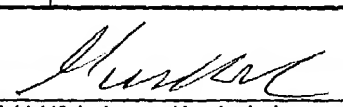
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JP	Pennisi, "Gene Linked to Commonest Cancer", Science 272: 1583-1584 (1996).		
CR	Perrimon et al., "Generating lineage-specific markers to study Drosophila development", Dev. Genet. 12:238-252 (1991).		
CS	Perrimon, N., "Serpentine proteins litter into the wingless and hedgehog fields", Cell, 86 (4): 513-516 (1996).		
CT	Phillips, R. et al., "The Drosophila segment polarity gene patched is involved in a position signalling mechanism in imaginal discs", Development, 110: 105-114 (1990).		
CU	Quinn, A. et al., "Chromosome 9 allele loss occurs in both basal and squamous cell carcinomas of the skin", J. Invest. Dermatology, 102: 300-303 (1994).		
CV	Quinn, A. et al., "Delineation of two distinct deleted regions on chromosome 9 in human non-melanoma skin cancers", Genes, Chromosomes & Cancers, 11:222-225 (1994).		
CW	Riddle, R. et al., "Sonic hedgehog mediates the polarizing activity of the ZPA", Cell, 75: 1401-1416 (1993).		
CX	Roelink, H. et al., "Floor plate and motor neuron induction by <i>vhh-1</i> , a vertebrate homolog of hedgehog expressed by the notochord", Cell, 76: 761-775 (1994).		
CY	Rogers, G. et al., "Patched gene mutation screening in patients with basal cell nevus syndrome using bidirectional dideoxy fingerprinting", J. Invest. Dermatol. Abstracts, 108(4): 598, # 364, (1997).		
CZ	Roush, W., "Hedgehog's patterning call is patched through, smoothly", Science, 274 (5291): 1304-1305 (1996).		
DA	Sanicola, M. et al., "Drawing a stripe in Drosophila imaginal disks: negative regulation of decapentaplegic and patched expression by engrailed", Genetics, 139 (2): 745-756 (1995).		
DB	Schuske, K. et al., "Patched overexpression causes loss of wingless expression in Drosophila embryos", Dev. Biol., 164 (1): 300-301 (1994).		
DC	Shilo, B., "Tumor suppressors. Dispatches from patched", Nature, 382 (6587): 115-116 (1996).		
DD	Simcox, A. et al., "Imaginal discs can be recovered from culture embryos mutant for the segment-polarity genes engrailed, naked and patched but not from wingless", Development, 107: 715-722 (1989).		
DE	Sisson, J. et al., "Costal2, a novel kinesin-related protein in the Hedgehog signaling pathway", Cell, 90 (2): 235-245 (1997).		
DF	Smith et al., "Genes Transferred by Retroviral Vectors into Normal and Mutant Myoblasts in Primary Cultures Are Expressed in Myotubes", Mol. Cell. Biol. 3268-3271 (1990).		
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		Stone, D. et al., "The tumor-suppressor gene patched encodes a candidate receptor for Sonic hedgehog", Nature, 384 (6605): 129-134 (1997).			
		Struhl, G. et al., "Hedgehog acts by distinct gradient and signal relay mechanisms to organize cell type d cell polarity in the Drosophila abdomen", Development, 124 (11): 2155-2165 (1997).			
	DI	Strutt, D. et al., "Ommatidial polarity in the Drosophila eye is determined by the direction of furrow progression and local interactions", Development, 121 (12): 4247-4256 (1995).			
	DJ	Strutt, D. et al., "Regulation of furrow progression in the Drosophila eye by cAMP-dependent protein kinase A", Nature, 373 (6516): 705-709 (1995).			
	DK	Tabata, T. et al., "The Drosophila hedgehog gene is expressed specifically in posterior compartment cells and is a target of engrailed regulation", Genes Dev., 6(12B): 2635-2645 (1992).			
	DL	Tabata, T. et al., "Hedgehog is a signaling protein with a key role in patterning Drosophila imaginal discs", Cell, 76: 89-102 (1994).			
	DM	Takabatake, T. et al., "Hedgehog and patched gene expression in adult ocular tissues", FEBS Letters, 410 (2-3): 485-489 (1997).			
	DN	Thummel, C. et al., "Vectors for Drosophila P-element mediated transformation and tissue culture transfection", Gene, 74: 445-446 (1988).			
	DO	Von Ohlen, T. et al., "Hedgehog signaling regulates transcription through cubitus interruptus, a sequence-specific DNA binding protein", Proc. Natl. Acad. Sci. USA, 94 (6): 2404-2409 (1997).			
	DP	Vorechovsky, I. et al., "Somatic mutations in the human homologue of Drosophila patched in primitive neuroectodermal tumors", Oncogene, 15 (3): 361-366 (1997).			
	DQ	Vortkamp, A., et al., "Regulation of rate of cartilage differentiation by Indian hedgehog and PTH-related protein", Science, 273 (5275): 613-622 (1996).			
	DR	Watson, J., Recombinant DNA, W. H. Freeman and Co., New York, 363, (1992).			
DS	Wicking, C. et al., "Fine genetic mapping of the gene for nevoid basal cell carcinoma syndrome", Genomics, 22: 505-511 (1994).				
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